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The method according to claim 1, wherein:

(a) step (a) of Claim 1 is carried out in a reaction medium which comprises an organic solvent, and optionally wherein said organic solvent comprises an organic base, and further optionally wherein said reactant comprises an acid anhydride, an acid chloride, a carboxylic acid or an N-acylimidazole, and further optionally wherein said reaction medium further comprises an acylation catalyst, and further optionally wherein said the reaction medium further comprises water;

- (b) said RNA comprises mRNA, rRNA or viral RNA;
- (c) \ said sample comprises a sample from a biological source;
- (d) said sample includes DNA;
- (e) sald substituent comprises a solid phase, and optionally wherein said solid phase comprises benzoyl chloride polymer bound (BCPB) beads, silica particles or particles of a glass, and further optionally wherein said solid phase is modified to introduce a reactive group which reactive group is capable of reacting with RNA to capture the RNA on the solid phase, and further optionally wherein said reactive group is introduced by modifying the solid phase with a bi-functional acid halide;
- (f) said substituent comprises a hydrophobic substituent, and optionally wherein said hydrophobic substituent comprises a substituent, OR, wherein R is selected from the group consisting of: C₁-C₃₆ alkyl; C₁-C₃₆ alkenyl; C₁-C₃₆ alkynyl; C₁-C₃₆ haloalkyl; C₁-C₃₆ aminoalkyl; C₁-C₃₆ alkoxyalkyl; C₁-C₃₆ alkylthioalkyl; C₁-C₃₆ alkoxyalkyl; C₁-C₃₆ haloalkoxyalkyl; C₁-C₃₆ arylalkyl; C₆-C₃₆ arylalkyl; C₆-C₃₆ arylalkyl; C₆-C₃₆ arylalkenyl; C₁-C₃₆ alkanoyl; C₁-C₃₆ alkenoyl; C₁-C₃₆ haloalkanoyl; C₁-C₃₆ haloalkanoyl; C₁-C₃₆ carboxyalkanoyl; C₁-C₃₆ carboxyalkynoyl; C₁-C₃₆ carboxyalkynoyl; C₁-C₃₆ carboxyalkynoyl; C₁-C₃₆ carboxyalkynoyl; C₁-C₃₆ carboxyalkynoyl;

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(g)

C₁-C₃₆ alkylaminoarylalkanoyl; C₁-C₃₆ alkoxycarbonyl; C₁-C₃₆ alkenyloxycarbonyl; C₁-C₃₆ alkylsulfonyl; C₆-C₃₆ arylalkanoyl; C₆- C_{36} arylalkenoyl; C_6 - C_{36} aryloxyalkanoyl; C_6 - C_{36} alkylarylalkanoyl; C₆-C₃₆ haloarylalkanoyl; C₆-C₃₆ aminoarylalkanoyl; C₁-C₃₆ alkylsilanyl; C₁-C₃₆ trialkylsilanyl and C₁₂-C₂₈ diarylphosphano; or a substituent R', wherein R' comprises C₁-C₃₆ alkyl; C₁-C₃₆ alkenyl; C_1 - C_{36} alkynyl; C_1 - C_{36} haloalkyl; C_1 - C_{36} aminoalkyl; halo; amino; C_1 - C_{36} alkylamino; C_6 - C_{36} aryl; C_1 - C_{36} alkylaryl or C_1 - C_{36} arylalkyl; said hydrophobic substituent of (f) comprises a C₄ to C₇ carbon chain or ring;

(h) wherein said reactant comprises butyric anhydride, pentanoic anhydride, hexanoic anhydride or benzoic anhydride;

(i) said proportion of 2'-OH positions bearing the substituent is at least 10%

said hydrophobic substituent of (f) comprises a C₈-C₁₂ carbon chain **(j)** or ring, and optionally wherein said proportion of 2'-OH positions bearing the substituent is in the range 1 to 10%;

said hydrophobia substituent of (f) comprises a C₁₂-C₃₆ carbon chain (k) or ring, and optionally wherein said proportion of 2'-OH positions bearing the substituent is up to 1%;

(l) said step (b) comprises contacting the treated sample from step (a) with a hydrophobic solid phase so as to bind the material containing the hydrophobic substituent and optionally washing the material bound to the solid phase, and optionally wherein said hydrophobic solid phase comprises hydrophobic particles, and further optionally wherein said method further comprises astep of eluting the material bound to the hydrophobic solid phase by treating with a detergent, a chaotrope or a solvent, by lowering the salt concentration or by cleaving the substituent from the 2'-OH position of the ribose rings; said step (b) comprises the further step of treating the treated sample (m) from step (a) with a lyotrophic salt to aggregate the material

containing the hydrophobic substituent as an RNA precipitate, and

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isolating the precipitate, and optionally wherein said lyotrophic salt comprises ammonium sulphate, an alkali metal chloride, magnesium chloride or calcium chloride; or

(n) said step (b) comprises treating the treated sample with a non-polar solvent to form a hydrophobic liquid phase which contains the material containing the hydrophobic substituent, and isolating the hydrophobic liquid phase, and optionally wherein said non-polar solvent comprises pentane, cyclohexane, toluene, benzene, light petroleum, xylene or hexane.

A kit for the preparative isolation of RNA comprising an oligo- or polynucleotide from a sample, which kit comprises:

- (i) a reaction system for modifying the RNA to form a modified oligoor poly-nucleotide in which a proportion of the 2'-OH positions of the ribose rings bear a substituent; and
- (ii) a separation system for preparing isolated RNA by separating material containing the substituent from the sample on the basis of a property of the substituent.

The kit according to Claim 3, wherein said reaction system comprises:

- (a) an organic solvent; and
- (b) a reactant capable of covalently modifying the 2'-OH position of the ribose rings of the RNA in the presence of the organic solvent, and optionally wherein:
 - (i) said organic solvent comprises an organia base;
 - (ii) said reactant comprises an acid anhydride, an acid chloride, a carboxylic acid or an N-acylimidazole;
 - (iii) said kit comprises an acylation catalyst;
 - (iv) said substituent comprises a solid phase, and optionally wherein said solid phase comprises benzoyl chloride

polymer bound (BCPB) beads, silica particles or particles of a glass; and optionally wherein:

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(c)

said substituent comprises a hydrophobic substituent, or more specifically wherein said hydrophobic substituent comprises a substituent, OR, wherein R comprises a moiety selected from the group consisting of: C₁-C₃₆ alkyl; C₁-C₃₆ alkenyl; C₁-C₃₆ alkynyl; C₁-C₃₆ haloalkyl; C₁-C₃₆ aminoalkyl; C₁-C₃₆ alkoxyalkyl; C₁-C₃₆ alkylthioalkyl; C₁-C₃₆ alkoxyalkoxyalkyl; C₁-C₃₆ haloalkoxyalkyl; $C_1 C_{36}$ aminoalkoxyalkyl; $C_6 - C_{36}$ aryl; $C_6 - C_{36}$ alkylaryl; $C_6 - C_{36}$ arylakyl; C₆-C₃₆ arylalkenyl; C₁-C₃₆ alkanoyl; C₁-C₃₆ alkenoyl; C₁-C₃₆ haloalkenoyl; C₁-C₃₆ haloalkanoyl; C₂-C₃₆ haloformylalkanoyl; C_1 - C_{36} C_1 - C_{36} aminoalkanoyl; C_1 - C_{36} azidoalkanoyl; C_1 - C_{36} carboxyalkanoyl; C1-C36 carboxyalkenoyl; C1-C36 carboxyalkynoyl; C₁-C₃₆ alkylaminoarylalkanoyl; C₁-C₃₆ alkoxycarbonyl; C₁-C₃₆ alkenyloxycarbonyl; C₁-C₃₆ alkylsulfonyl; C₆-C₃₆ arylalkanoyl; C₆-C₃₆ arylalkenoyl; C₆-C₃₆ aryloxyalkanoyl; C₆-C₃₆ alkylarylalkanoyl; C₆-C₃₆ haloarylalkanoyl; C₆-C₃₆ aminoarylalkanoyl; C₁-C₃₆ alkylsilanyl; C1-C36 trialkylsilanyl and C12-C28 diarylphosphano; or a substituent R', wherein R' domprises C₁-C₃₆ alkyl; C₁-C₃₆ alkenyl; C₁-C₃₆ alkynyl; C₁-C₃₆ haloalkyi; C₁-C₃₆ aminoalkyl; halo; amino; C_1 - C_{36} alkylamino; C_6 - C_{36} aryl; C_1 - C_{36} alkylaryl or C_1 - C_{36} arylalkyl.

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The kit according to claim 4, wherein:

- (a) said hydrophobic substituent comprises a C₄ to C₇ carbon chain or ring;
- (b) said reactant comprises butyric anhydride, pentanoic anhydride, hexanoic anhydride or benzoic anhydride;
 - (c) said proportion of 2'-OH positions bearing the substituent is at least 10%;

containing the substituent from the sample on the basis of a property

said hydrophobic substituent comprises a C₈-C₁₂ carbon chain or 9 ring; said proportion of 2'-OH positions bearing the substituent is 10 (e) selected from any one integer from 1 to 10% inclusive; 11 said hydrophobic substituent comprises a C₁₂-C₃₆ carbon chain or (f) 13 ring; said proportion of 2'-OH positions bearing the substituent is up to 14 (g) 15 1%; said separation system comprises a hydrophobic solid phase for 16 (h) binding the material containing the substituent, and optionally 17 wherein said hydrophobic solid phase comprises hydrophobic 18 particles, and further optionally wherein said separation system 19 further comprises an elution medium for eluting RNA bound to the 20 21 hydrophobic solid phase; 22 said separation system comprises a lyotrophic salt for aggregating (i) 23 the material containing the hydrophobic substituent; or 24 said separation system comprises a non-polar solvent for forming a (j) 25. hydrophobic liquid phase which contains the material containing the 26 hydrophobic substituent. (၁၃ A preparative device for isolating RNA comprising an oligo-or polynucleotide from a sample from a subject, which device comprises: 2 a means for extracting the sample from the subject; 3 (i) a reaction system for modifying RNA in the sample to form a (ii) 5 modified oligo- or poly-nucleotide in which a proportion of the 2'-OH positions of the ribose rings bear a substituent; and 6 7 (iii) a separation system for preparing isolated RNA by separating material

of the substituent.

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The device according to claim 6, wherein: said means for extracting the sample from the subject comprises a (a) 2 syringe needle; 3 said substituent comprises a solid phase, and optionally wherein the (b) solid phase comprises a membrane, a particle, a bead, a filter, a fibre, a gel, a strip, a matrix, a resin, a capillary or the walls of a vessel; said sample comprises biological material; or (c) said device further comprises a filter for removing red and/or white (d) blood cells.

> The Commissioner is hereby authorized to charge any additional fees or credit any overpayment to the Credit Card listed on PTO Form 2038, filed with this application.

> > Respectfully submitted,

26 OCT, 2001 By:

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